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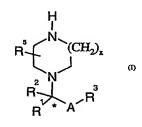
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(54) Title: PIPERAZINE DERIVATIVES WHICH EXHIBIT ACTIVITY AS SEROTONIN AND NORADRENALINE RE-UP-ATKE INHIBITORS



(57) Abstract: A compound of formula (I), wherein R¹ is H; R² is aryl, het, C_{3.8}cycloalkyl, C₁₋₆alkyl, (CH₂)₂aryl or R⁴, wherein each of the cycloalkyl, aryl, het and R⁴ groups is optionally substituted by at least one substituent independently selected from C₁₋₆alkyl, C₁₋₆alkoxy, OH, halo, CF_3 , OCF_3 , $OCHF_2$, $O(CH_2)_yCF_3$, CN, $CONH_2$, $CON(H)C_{1-6}alkyl$, $CON(C_{1-6}alkyl)_2$, hy $droxy-C_{1\text{-}6}alkyl,\ C_{1\text{-}4}alkoxy-C_{1\text{-}6}alkyl,\ C_{1\text{-}4}alkoxy-C_{1\text{-}4}alkoxy,\ SCF_3,\ C_{1\text{-}6}alkylSO_2,\ C_{1\text{-}4}alkyl-S-C_{1\text{-}6}alkylSO_2,\ C_{1\text{-}4}alkyl-S-C_{1\text{-}6}alkyl-S-C_{1\text{-$ 4alkyl, C14alkyl-S-, C14alkyINR10R11 and NR10R11; or R1 and R2, together with the carbon atom to which they are bound, form a 5- or 6-membered carbocycle or a 5- or 6-membered heterocycle containing at least one N, O or S heteroatom; R3 is aryl, het or R4, each optionally substituted by at least one substituent independently selected from C₁₋₆alkyl, C₁₋₆alkoxy, het, OH, halo, CF₃, OCF₃, OCHF₂, O(CH₂)₇CF₃, CN, CONH₂, CON(H)C₁₋₆alkyl, CON(C₁₋₆alkyl)₂, hydroxy-C₁.

 $_{6}$ alkyl, C_{1-4} alkoxy- C_{1-6} alkyl, C_{1-4} alkoxy- C_{1-4} alkoxy, SCF₃, C_{1-6} alkylSO₂, C_{1-4} alkyl-S- C_{1-4} alkyl, C_{1-4} alkyl-S-, C_{1-4} alkylNR 10 R 11 and NR¹⁰R¹¹; R⁴ is a phenyl group fused to a 5- or 6 membered carbocycle, or a phenyl group fused to a 5- or 6-membered heterocycle containing at least one N, O or S heteroatom; R⁵ is H or C_{1.6} alkyl; R¹⁰ and R¹¹ are the same or different and are independently H or C₁₋₄alkyl; A is a C₁₋₃alkylene chain which is optionally substituted by OH, C₁₋₄alkyl or C₁₋₄alkoxy; x is an integer from 1 to 3; y is 1 or 2; z is an integer from 1 to 3; aryl is phenyl, naphthyl, anthracyl or phenanthryl; and het is an aromatic or non-aromatic 4-, 5- or 6-membered heterocycle which contains at least one N, O or S heteroatom, optionally fused to a 5- or 6-membered carbocycle or a second 4-, 5- or 6-membered heterocycle which contains at least one N, O or S heteroatom.

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